Supplemental Figure 1. dGTP induces SAMHD1 tetramerization. A. SAMHD1 ( $10~\mu M$ ) was preincubated in the absence (A), or presence of 1 mM (B) or 4 mM (C) dGTP. The mixtures were injected into an analytical gel filtration column and separated at a flow rate of 0.8 mL/min. Elution profile (A280 nm) was recorded. The elution positions of tetramer and dimer/monomer peaks are indicated.

Supplemental Figure 2. SAMHD1 tetramer and dimer/monomer interconvert in a dGTP-dependent manner. A. A mixture of SAMHD1 (25  $\mu$ M) and dGTP (45  $\mu$ M) was injected to an analytical gel filtration column at a flow rate of 0.8 mL/min. Elution profiles (A254 nm, red trace, and A280 nm, blue trace) were recorded. Elution fractions corresponding to tetramer (1) and dimer/monomer (2) were collected, concentrated, and analyzed by SDS-PAGE and visualized by Coomassie Blue staining. The fraction containing tetramer (B) and dimer/monomer (C) from A were subjected to analytical gel filtration column chromatography. Elution profiles as revealed by fluorescence trace (excitation at 282 nm and emission at 313 nm) were recorded. D. The same fraction containing tetramers (1 in (A)) was mixed with 4 mM dGTP prior to gel filtration analysis. E. The fraction containing SAMHD1 dimer/monomer (2 in (A)) was incubated with 4 mM dGTP and then subjected to gel filtration analysis. F. The elution fraction containing tetramer shown as (1) in (A) was subjected to dNTPase assays a reaction buffer without or with dGTP (25  $\mu$ M) added. The dA nucleoside product was quantified by HPLC. G. The elution fraction containing dimer/monomer fraction shown as (2) in (A) was subjected to dNTPase assays performed as in panel F.







